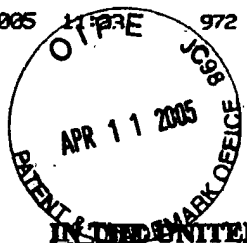


10.APR.2005 11:33E 972 04 6361004

NO.008 P.6

**IN THE UNITED STATES PATENT AND TRADEMARK OFFICE**

In re Application of:

LAHAV et al

Serial No.: 10/018,992

Filed: February 19, 2003

For: STABLE BENZIMIDAZOLE
FORMULATION

Examiner: OH, SIMON J

Commissioner for Patents
PO Box 1450
Alexandria, VA 22313-1450

Group Art Unit: 1615

Attorney
Docket: 24305
Previously: D01/200**DECLARATION UNDER 37 C.F.R. § 1.31(a)**

I, Dr. Valerie Azoulay, declare as follows:

1) I am Dr. Valerie Azoulay who is an inventor named in the above-identified subject invention.

2) I understand that the pending claims, claims 40 to 90, are rejected under 35 U.S.C. section § 103(a) as allegedly unpatentable over Dietrich et al. (WO 99/27917; hereinafter "Dietrich") in view of Lundberg et al. I also understand that Dietrich was published on June 10, 1999.

3) The priority date of the above-identified application is June 22, 1999, which is well within one year of the publication date of Dietrich.

4) Moreover, the subject invention is directed to a stable formulation for a benzimidazole derivative by having a single neutralized enteric coating layer. This

858 459 0698

10.APR.2005 11:23

972 04 6364004

NO.008 P.7

Declaration of Valerie Azoulay
U.S. App. Ser. No. 10/018,992
Page 2

invention was conceived and reduced to practice prior to the publication date of Dietrich. Therefore, Dietrich is unsuitable as prior art. In support of this position attached hereto are exhibits, provided both in the original language and fully translated into English.

5) More specifically, an experiment involving neutralizing an enteric coating by applying an alkaline compound, was performed by the Applicant on January 19, 1999. This experiment featured a benzimidazole compound, omeprazole (internally coded as "ODYSSEUS"), formulated into tablet cores, which were then coated with an enteric coating. The results showed that the enteric polymer was neutralized with an alkaline compound, in this particular case, sodium hydroxide. The Certificate of Analysis for this experiment (Exhibit A) is dated January, 27, 1999. The Certificate indicates that the tablets were coated with an enteric coating and brought to pH of 5.8. (See line 3 of Exhibit A). The instructions for this experiment indicates that a 1 mol/l NaOH solution (the neutralizing compound) was mixed with the coating suspension to achieve this pH. (See section 8 of Exhibit B).

6) Further experiments were performed from January 19, 1999 to March 15, 1999. Many of these experiments exemplify the subject invention as claimed. In these experiments other polymers were neutralized. The resulting enteric coating from several of these experiments had pH's of 7.39, 7.48 and 8.03. (See for example, sections 1.4.7 and 1.4.9 of Exhibit C; and section 1.4.7 of Exhibit D, respectively).

858 459 0698

PAGE 3/10

APR-10-05 4:53AM;

858 459 0698;

IT BY: LAW OFFICE OF DAVID SPOLTER;

10.APR.2005 11:24

972 04 6364004

NO.008 P.8

Declaration of Valerie Azoulay
U.S. App. Ser. No. 10/018,992
Page 3

I declare that all statements made herein of my knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements are made with the knowledge that willfully false statements are punishable by fine or imprisonment under 18 U.S.C. Section 1001 and that any such statement may jeopardize the validity of the subject application or any patent issued thereon.

Azoulay
Dr. Valerie Azoulay

10/4/05
Date



IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of:

LAHAV et al

Serial No.: 10/018,992

Filed: February 19, 2003

**For: STABLE BENZIMIDAZOLE
FORMULATION**

Examiner: OH, SIMON J

Commissioner for Patents
P.O. Box 1450
Alexandria, VA 22313-1450

~~~~~

**Group Art Unit: 1615**

**Attorney**  
**Docket: 24305**  
**Previously: D01/200**

## CERTIFICATION

Sir:

I, the undersigned Sol Sheinbein, hereby declare that I am well acquainted with the Hebrew and English languages and that the document attached hereto is a correct English translation of the original document drawn up in the Hebrew language.

In witness whereof I certify the correctness of the said translation by my signature  
on this 11<sup>th</sup> day of April 2005.

Respectfully submitted,

Ad. Hake

**Sol Sheinbein**  
**Registration No. 25,457**

**Date: April 11, 2005**



10 APR 2005 11:24

972 04 6364004

NO. 008 P. 10

## TRANSLATION INTO ENGLISH OF EXHIBIT A

*I checked the translation and found it accurate*  
Shirley Wheeler, RPh.  
Dexcon Ltd.

Dexcon Ltd.

R&amp;D DEPARTMENT

## CERTIFICATE OF ANALYSIS NO' 1854

Product ODYSSEUS 20mgBatch No. 221298/C1/C2 quantity \_\_\_\_\_ stage of production after enteric coating  
(697mg) (audimet L90D pH 5.8)Received by Valerie date 27.1.99 time 8:00

| Test                                              | method/expected range | results | remarks   |
|---------------------------------------------------|-----------------------|---------|-----------|
| Dissolution 2hrs in gastric fluid and then pH 6.8 |                       |         | check 3/3 |

## DISSOLUTION

pH: 2 hr gastric fluid buf. 6.8 vol: 900 ml equipment: paddle r.p.m: 100samples times: 1.2.3 - 221298/C1  
4.5.6 - 221298/C2

| Vessel No. | 1      | 2      | 3      | 4      | 5      | 6      |
|------------|--------|--------|--------|--------|--------|--------|
| Tab Weight | 0.7158 | 0.7212 | 0.6918 | 0.7374 | 0.7318 | 0.7496 |

## ATTACH DISSOLUTION PROFILE

Result file

Laboratory number: 5186 analyst's signature: \_\_\_\_\_

Notebook: Glucomin forte no. 1 Date: 31.01.99

Signature of responsible R&amp;D group \_\_\_\_\_

signature of R&amp;D laboratory manager \_\_\_\_\_

10.APR.2005 11:24

972 04 6364004

NO.008

P.12

GHB (PM)

7. פרום תהליך העבודה:

7.1 הוצא טבליות לא מצופות במשקל  $2.000 \text{ g}$  לשקית מסומנת בהתאם, סלל ציון מצב הטבליות (UNCOATED).

תאריך: 25/1/99 שם וחתימת העובד: א-א

7.2 שיקול את הטבליות הפעילות והבלתי פעילות.  
סה"כ צריך להיות בחד כ- 2.000 קג

| טבליות פעילות  | טבליות בלתי פעילות |                 |
|----------------|--------------------|-----------------|
|                |                    | משקל ברוטו      |
|                |                    | משקל טרו        |
| <u>2000</u> קג | —                  | משקל טבליות נטו |

תאריך: 25/1/99 שם וחתימת העובד: א-א

7.3 מפרט הטבליות הפעילות והבלתי פעילות.

| טבליות פעילות | טבליות בלתי פעילות |                                 |
|---------------|--------------------|---------------------------------|
|               |                    | מידות טבליות (אורך, רוחב, קוטר) |
|               |                    | עובי                            |
| <u>597mg</u>  | —                  | משקל ממוצע של טבליות (ס)        |
|               |                    | טבליות                          |

תאריך: 25/1/99 שם וחתימת העובד: א-א

8. תרכב תרופה המפוי:

3333 tablets

| תאריך          | חתימה      | כמות (קג)       | חומר גלם                       | Mg/Tab     | הערות |
|----------------|------------|-----------------|--------------------------------|------------|-------|
| <u>25/1/99</u> | <u>א-א</u> | <u>2kg</u>      | <u>ONDUSSELS</u>               | <u>600</u> |       |
| <u>25/1/99</u> | <u>א-א</u> | <u>1.050 קג</u> | <u>Water</u>                   |            |       |
| <u>25/1/99</u> | <u>א-א</u> | <u>0.700 קג</u> | <u>Calcium chloride 1mol/l</u> |            |       |
| <u>25/1/99</u> | <u>א-א</u> | <u>0.700 קג</u> | <u>Eudragit L30</u>            |            |       |
| <u>25/1/99</u> | <u>א-א</u> | <u>0.021 קג</u> | <u>Calcium citrate</u>         |            |       |
| <u>25/1/99</u> | <u>א-א</u> | <u>0.064 קג</u> | <u>Talc</u>                    |            |       |
|                |            |                 |                                | <u>680</u> |       |
|                |            |                 |                                | <u>700</u> |       |
|                |            |                 |                                | <u>720</u> |       |
|                |            |                 |                                |            | סה"כ  |

10. APR. 2005 11:25

972 84 6364804

**TRANSLATION INTO ENGLISH OF EXHIBIT B**

*I checked the translation and found it accurate*  
*Stefy Weber, RPh*  
*Degea Ltd*

**Instructions for tablet coating experiments: ODYSSEUS TABLET****Batch number: 221298/C product number: 115123 date: 24.1.99 quantity of tablets: 2kg**

1. List of necessary equipment for coating:
- |                                |                      |
|--------------------------------|----------------------|
| main equipment                 | supporting equipment |
| - THAI COATER/ AGGELA-COTA-160 | plastic boxes        |
| - Mixer                        | polyethylene bags    |
| - FLOWTAB/SPRAYTAB             | cars-blockers        |
| - Differential balances        |                      |

2. I confirm that the area, dishes and machines for the preparations of the coating solutions are clean. The raw material that is needed for the coating is complete and approved and the coating instructions are complete. I approve starting the preparation of the solutions.

Date: 25.1.99 name and signature of researcher/QA \_\_\_\_\_

3. I approve the starting of the coating stage.

Date: 25.1.99 name and signature of researcher/QA \_\_\_\_\_

4. remarks: coating with endragit L30D the pH of the solution: 5.75-5.8  
coat in a low temperature 40 deg.C - low spray rate with double amount of water.

Date: 24.1.99 name and signature of researcher/QA \_\_\_\_\_

5. I confirm that the area, dishes and machines are clean. The necessary tablets are prepared and approved for coating. The coating dispersion is prepared according to the instructions. I approve starting the coating.

In addition I am releasing the tablets from "quarantine".

Date: 25.1.99 name and signature of researcher/QA \_\_\_\_\_

6. Mark the equipment with the product name, batch number and the necessary quantity.

Date: 25.1.99 name and signature of employee \_\_\_\_\_



10.APR.2005 11:25

972 04 6364004

NO.008 P.14

## 7. details of the working procedure:

7.1 take the uncoated tablets in a weight of 2.000 kg to marked bags accordingly that indicate the status of the tablets (uncoated).

Date: 25.1.99 name and signature of employee \_\_\_\_\_

7.2 weigh the active and inactive (placebo) tablets.  
a pan should contain a total weight of about 2.000 kg

|             | Inactive tablets (placebo) | active tablets |
|-------------|----------------------------|----------------|
| Brut weight |                            |                |
| Tara weight |                            |                |
| Net weight  | -                          | 2.000 kg       |

Date: 25.1.99 name and signature of employee \_\_\_\_\_

## 7.3 specification for active and inactive tablets

|                                           | Inactive tablets (placebo) | active tablets |
|-------------------------------------------|----------------------------|----------------|
| Tablet dimension (length, wide, diameter) |                            |                |
| Thickness                                 |                            |                |
| Tablet mean weight (50 tablets)           |                            | 597 mg         |

Date: 25.1.99 name and signature of employee \_\_\_\_\_

## 8. composition of coating dispersion:

3333 tablets

| date    | signature | quantity (kg) | raw material             | mg/tab | remarks |
|---------|-----------|---------------|--------------------------|--------|---------|
| 25.1.99 |           | 2 kg          | ODYSSSEUS                | 600    |         |
| 25.1.99 |           | 1.050 kg      | water                    |        |         |
| 25.1.99 |           | 0.700 kg      | solution NaOH 1mol/liter |        |         |
| 25.1.99 |           | 0.700 kg      | Budragit L30D            |        |         |
| 25.1.99 |           | 0.021 kg      | triethyl citrate         |        |         |
| 25.1.99 |           | 0.064 kg      | talc                     |        |         |
|         |           | total         |                          | 680    |         |
|         |           |               |                          | 700    |         |
|         |           |               |                          | 720    |         |

10.APR.2005 11:25

972 04 6364004

NO. 008 P. 15

**Exh C (PA)**

OMEPRAZOLE 20 mg

מידע סודי לשימוש פנימי בלבד

"OM1232A1"

22/3/99 תאריך  
 שם העבד  
 חתימה

1.3.4 השלם עם מים מטופלים עד פס הבקבוק של 1 ליטר, המסך לערבב בעזרת STIRRER + גלטה MAGNET עד שהתמיסה מגיעה לטמפרטורה החדר.

## 1.4 הכנת תרחיף ציפוי אנטרי

1.4.1 שמן חציר בתווית "מזרז בנחליך".

22/3/99 תאריך  
 שם העבד  
 חתימה

1.4.2 לתוך דלי הפלסטיק של 5 ליטר המסומן כמזרז שקולה שלו. תאריך: 22/3/99

| משקל     | חומר        | שם וחתימת עובד 1 | שם וחתימת עובד 2 |
|----------|-------------|------------------|------------------|
| 0.372 KG | מים מטופלים | MA               | MA               |

1.4.3 חוסף תוך כדי עירבוב במערבל עד לחמשה, כמות שקולה שלו. תאריך: 22/3/99

| משקל     | חומר             | שם וחתימת עובד 1 | שם וחתימת עובד 2 |
|----------|------------------|------------------|------------------|
| 0.076 KG | TRIETHYL CITRATE | MA               | MA               |

1.4.4 חוסף תוך כדי עירבוב במערבל עד לחמשה, כמות שקולה שלו. תאריך: 22/3/99

| משקל     | חומר                   | שם וחתימת עובד 1 | שם וחתימת עובד 2 |
|----------|------------------------|------------------|------------------|
| 0.009 KG | SODIUM LAURYL SULPHATE | MA               | MA               |

1.4.5 חוסף תוך כדי עירבוב, כמות שקולה שלו. תאריך: 22/3/99

| משקל     | חומר  | שם וחתימת עובד 1 | שם וחתימת עובד 2 |
|----------|-------|------------------|------------------|
| 0.271 KG | AQOAT | MA               | MA               |
| 0.135 KG | TALC  | MA               | MA               |

10.APR.2005 11:25

972 04 6364004

NO. 008 P. 16

**Exh C (P2)**

OMEPRazole 20 mg

מידע סודי לשימוש פנימי בלבד

"OM1232A1"

תאריך: 22/3/99

1.4.6 חוסף ועד כדי ערבוב:

| שם וחתימת עובד 2 | שם וחתימת עובד 3 | חומר                             | משקל     |
|------------------|------------------|----------------------------------|----------|
| <i>Chouley</i>   | <i>W. O. K.</i>  | SOLUTION SODIUM<br>HYDROXYDE 1 M | 0.360 KG |

המשך לערבב 30 דקות.

22/3/99

תאריך

10<sup>00</sup>

זמן ותחלה:

10<sup>30</sup>

זמן סיום:

22/3/99

שם העובד

22/3/99

חתימה

1.4.7 בדיקת PH של חתרוניף בעזרת PH METER.

9.39 = PH

22/3/99

תאריך

22/3/99

שם העובד

22/3/99

חתימה

1.4.8 במקרה ש-PH לא בתוך הטווח של 8-10 חוסף

SOLUTION SODIUM HYDROXYDE 1 M, בדרגות של 20gr

וערבב 15 דקות אחרי כל תוספת.

בדיקת PH אחרי כל תוספת.

22/3/99

תאריך

22/3/99

שם העובד

22/3/99

חתימה

1.4.9 לבצע את המעלות מסעיף 1.4.7 עד קבלת PH רצוי.

22/3/99

תאריך

22/3/99

שם העובד

22/3/99

חתימה

1.4.10 כמויות סופיות של 1 M SOLUTION SODIUM HYDROXYDE

0.020 kg

= שחוסף

1.03

= PH סופי

22/3/99

תאריך

22/3/99

שם העובד

22/3/99

חתימה

1.5 אישור הטבליות לציפוי

22/3/99

תאריך

22/3/99

שם העובד

22/3/99

חתימה

1.5.1 חתק מידף מעקב ביצוע דחיסה" את המשקל הממוצע

לפני ציפוי של טבליות 529 מ"ג.

10.APR.2005 11:26

972 04 6364004

NO.008 P.17

**TRANSLATION INTO ENGLISH OF EXHIBIT C**

*I checked the translation and found it accurate*  
*Stefy Weber, Adv.*  
*Dagyn Ltd.*

**Confidential Information For internal use only**

**1.4 Preparation of enteric coating suspension**

**1.4.1 label the equipment with the sticker: "product in process"**

date 22.3.99

name of employee \_\_\_\_\_

signature \_\_\_\_\_

**1.4.2 Introduce a measured amount of the following into a 5 liters container of stainless- still**

date: 22.3.99

| Weight   | Raw material   | Name & signature of 1 <sup>st</sup> employee | Name & signature of 2 <sup>nd</sup> employee |
|----------|----------------|----------------------------------------------|----------------------------------------------|
| 3.372 KG | Purified water |                                              |                                              |

**1.4.3 add while mixing in the mixer until dissolving a measured amount of:**

date: 22.3.99

| Weight   | Raw material     | Name & signature of 1 <sup>st</sup> employee | Name & signature of 2 <sup>nd</sup> employee |
|----------|------------------|----------------------------------------------|----------------------------------------------|
| 0.076 KG | TRIETHYL CITRATE |                                              |                                              |

**1.4.4 add while mixing in the mixer until dissolving a measured amount of:**

date: 22.3.99

| Weight   | Raw material           | Name & signature of 1 <sup>st</sup> employee | Name & signature of 2 <sup>nd</sup> employee |
|----------|------------------------|----------------------------------------------|----------------------------------------------|
| 0.009 KG | SODIUM LAURYL SULPHATE |                                              |                                              |

**1.4.5 add while mixing a measured amount of:**

date: 22.3.99

| Weight   | Raw material | Name & signature of 1 <sup>st</sup> employee | Name & signature of 2 <sup>nd</sup> employee |
|----------|--------------|----------------------------------------------|----------------------------------------------|
| 0.271 KG | AQOAT        |                                              |                                              |
| 0.135 KG | TALC         |                                              |                                              |

**1.4.6 add while mixing:**

date: 22.3.99

| Weight   | Raw material                 | Name & signature of 1 <sup>st</sup> employee | Name & signature of 2 <sup>nd</sup> employee |
|----------|------------------------------|----------------------------------------------|----------------------------------------------|
| 0.360 KG | SOLUTION SODIUM HYDROXIDE 1M |                                              |                                              |

10.APR.2005 11:26

972 04 6364004

NO.008 P.18

**Continue mixing 30 minutes****Beginning time: 10:00****End time: 10:30**date 22.3.99

name of employee \_\_\_\_\_

signature \_\_\_\_\_

**1.4.7 check pH of the suspension by a pH meter****pH= 7.39**date 22.3.99

name of employee \_\_\_\_\_

signature \_\_\_\_\_

**1.4.8 in case the pH is not in the limits of 8-10, add SOLUTION SODIUM HYDROXIDE 1 M in portions of 20 gr and mix 15 minutes after every addition.****Check pH after every addition.**date 22.3.99

name of employee \_\_\_\_\_

signature \_\_\_\_\_

**1.4.9 perform the actions starting in section 1.4.7 until obtaining the desired pH.**date 22.3.99

name of employee \_\_\_\_\_

signature \_\_\_\_\_

**1.4.9 final quantity of SOLUTION SODIUM HYDROXIDE 1M****added = 0.020 kg****final pH = 8.03**

10.APR.2005 11:26

972 04 6364004

NO.008

P.19

החומר (PA)

מועד סחי לשימוש פנימי בלבד

OMSPRAZOLE 20 mg

"OM1312A1"

1.3.4. חשלום עם מים מטופלים עד פס המקבוק של 1 ליטר, המטוך לערבב בעזרת STIRRER + פלטה MAGNET עד שהתמיסה מגיעה לסמך תחת.

4/6/99 תאריך  
1-0/16 שם חומר  
חתימה

1.4. הכנת תרחיק ציפוי אנטרי

1.4.1. סמן תצוה כמות "מוצר בתחילת".

4/6/99 תאריך  
1-0/16 שם חומר  
חתימה

1.4.2. לעד זלי הפלבים של 5 ליטר המסומן חכס במות שקולה שלו

| משקל     | חומר        | שם וחתימת עובד 1 | שם וחתימת עובד 2 |
|----------|-------------|------------------|------------------|
| 2.550 KG | מים מטופלים | 1-0/16           | אפולו            |

1.4.3. חוסף תוך כדי עירוב במערבל עד לחמסה, כמות שקולה שלו

| משקל     | חומר             | שם וחתימת עובד 1 | שם וחתימת עובד 2 |
|----------|------------------|------------------|------------------|
| 0.058 KG | TRIETHYL CITRATE | 1-0/16           | אפולו            |

1.4.4. חוסף תוך כדי עירוב במערבל עד לחמסה, כמות שקולה שלו

| משקל     | חומר                   | שם וחתימת עובד 1 | שם וחתימת עובד 2 |
|----------|------------------------|------------------|------------------|
| 0.006 KG | SODIUM LAURYL SULPHATE | 1-0/16           | אפולו            |

1.4.5. חוסף תוך כדי עירוב, כמות שקולה שלו

| משקל     | חומר  | שם וחתימת עובד 1 | שם וחתימת עובד 2 |
|----------|-------|------------------|------------------|
| 0.205 KG | AQOAT | 1-0/16           | אפולו            |
| 0.104 KG | TALC  | 1-0/16           | אפולו            |

18.APR.2005 11:26

972 04 6364004

NO. 008 P. 20

EXH (P2)

OMEPRAZOLE 20 mg

מידע סודי לשימוש פנימי בלבד

"OM(312A1)"

4/6/99

תאריך:

1.4.6 תוסף ועד כדי ערבוב:

| שם ותחומת עובד 1 | שם ותחומת עובד 2 | חומר                          | משקל     |
|------------------|------------------|-------------------------------|----------|
| יפלט             | 1-04             | SOLUTION SODIUM HYDROXIDE 1 M | 0.275 KG |

4/6/99

תאריך:

שם העובד

חתימה

640

ומן התחלת:

710

ומן סיום:

המשך לערבב 20 דקות.

1.4.7 בדיקת PH של התרחיף בעזרת PH METER,

3.48 = PH

4/6/99

תאריך:

שם העובד

חתימה

1.4.8 במקרה ש-PH לא בועד הגבולות של 7-9 תוסף

SOLUTION SODIUM HYDROXYDE 1 M, בדרגות של 20

וערבב 15 דקות אחרי כל תוספת.

בדיקת PH אחרי כל תוספת.

4.6.99

1.4.9 לבצע את הפעולות מסעיף 1.4.7 עד קבלת PH רצוי.

תאריך:

שם העובד

חתימה

1.4.10 בדיקת סופיות של SOLUTION SODIUM HYDROXYDE 1 M

שחוסף =

PH סופי =

תאריך:

שם העובד

חתימה

1.5 אישור הטבליות לצינור

4/6/99

תאריך:

שם העובד

חתימה

1.5.1 העתק מ"דף מעקב ביצוע ז'חיסור" את המשקל הממוצע

לפני ציפוי של טבליות 252 מ"ג.

10. APR. 2005 11:27

972 04 6364004

NO. 008

P. 21

**TRANSLATION INTO ENGLISH OF EXHIBIT D**

*I checked the translation and found it accurate*  
*Shirley Weiser, RPh*  
*Deegen Ltd*

**Confidential Information For Internal Use Only**

**1.4 Preparation of enteric coating suspension**

**1.4.1 Label the equipment with the sticker: "product in process"**

date 4.6.99

name of employee \_\_\_\_\_

signature \_\_\_\_\_

**1.4.2 Introduce a measured amount of the following into a 5 liters container of stainless- still**

date: 4.6.99

| Weight   | Raw material   | Name & signature of 1 <sup>st</sup> employee | Name & signature of 2 <sup>nd</sup> employee |
|----------|----------------|----------------------------------------------|----------------------------------------------|
| 2.550 KG | Purified water |                                              |                                              |

**1.4.3 add while mixing in the mixer until dissolving a measured amount of:**

date: 4.6.99

| Weight   | Raw material     | Name & signature of 1 <sup>st</sup> employee | Name & signature of 2 <sup>nd</sup> employee |
|----------|------------------|----------------------------------------------|----------------------------------------------|
| 0.058 KG | TRIETHYL CITRATE |                                              |                                              |

**1.4.4 add while mixing in the mixer until dissolving a measured amount of:**

date: 4.6.99

| Weight   | Raw material           | Name & signature of 1 <sup>st</sup> employee | Name & signature of 2 <sup>nd</sup> employee |
|----------|------------------------|----------------------------------------------|----------------------------------------------|
| 0.006 KG | SODIUM LAURYL SULPHATE |                                              |                                              |

**1.4.5 add while mixing a measured amount of:**

date: 4.6.99

| Weight   | Raw material | Name & signature of 1 <sup>st</sup> employee | Name & signature of 2 <sup>nd</sup> employee |
|----------|--------------|----------------------------------------------|----------------------------------------------|
| 0.205 KG | AQOAT        |                                              |                                              |
| 0.104 KG | TALC         |                                              |                                              |

**1.4.6 add while mixing:**

date: 4.6.99

| Weight   | Raw material                 | Name & signature of 1 <sup>st</sup> employee | Name & signature of 2 <sup>nd</sup> employee |
|----------|------------------------------|----------------------------------------------|----------------------------------------------|
| 0.275 KG | SOLUTION SODIUM HYDROXIDE 1M |                                              |                                              |



10.APR.2005 11:27

972 04 6364024

NO.008 P.22

Continue mixing 30 minutes

Beginning time: 06:40End time: 07:10date 4.6.99

name of employee \_\_\_\_\_

signature \_\_\_\_\_

1.4.7 check pH of the suspension by a pH meter

pH= 7.48date 4.6.99 name of

employee \_\_\_\_\_

signature \_\_\_\_\_

~~1.4.8 in case the pH is not in the limits of 7.9, add SOLUTION SODIUM  
HYDROXIDE 1 M in portions of 20 gr and mix 15 minutes after every  
addition.~~

~~Check pH after every addition.~~

date \_\_\_\_\_

name of employee \_\_\_\_\_

signature \_\_\_\_\_

1.4.9 ~~perform the actions starting in section 1.4.7 until obtaining the desired pH.~~

date \_\_\_\_\_

name of employee \_\_\_\_\_

signature \_\_\_\_\_

1.4.10 ~~final quantity of SOLUTION SODIUM HYDROXIDE 1M~~~~added =~~~~final pH =~~

date \_\_\_\_\_

name of employee \_\_\_\_\_

signature \_\_\_\_\_

1.5 **Appovement Of Tablets For Coating**1.5.1 Copy from the "Follow-up of compress document" the average weight  
before coating of the tablet 252 mgdate 4/6/99

name of employee \_\_\_\_\_

signature \_\_\_\_\_

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